

cell, and wherein expression of the selectable marker is operatively linked to expression of a Sox gene;

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(ii) culturing the pluripotent cell *in vitro* to induce differentiation of the pluripotent cell into a neural progenitor cell or into a mixture of cells including neural progenitor cells, or to induce preferential survival, in a mixed culture of cells, of neural progenitor cells; and

(iii) selecting for neural progenitor cells according to differential expression of the selectable marker introduced in step (i).

44. (Twice Amended) A method according to Claim 42 wherein the pluripotent cell is selected from embryonic stem (ES) cells, embryonic germ (EG) cells, embryonic carcinoma (EC) cells, a primary culture of fetal cells, a primary culture of post-natal cells, and a primary culture of adult cells.

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45. (Twice Amended) A method according to Claim 42 comprising genetically modifying pluripotent cells by deleting, mutating, substituting or adding genes in said pluripotent cells in order (i) to assay gene function in neural progenitor cells, or (ii) to render selected cells more suitable for transplantation, or both.

46. (Twice Amended) A method according to Claim 42 further comprising:

(iv) introducing into the pluripotent cell a second selectable marker that is differentially expressed in cells of a selected sub-lineage compared with its expression in other cells, wherein cells of the selected sub-lineage are formed by differentiation of neural progenitor cells; and

(v) when a culture of neural progenitor cells has been obtained, allowing or inducing differentiation of the cells and selecting for cells that express the second selectable marker.

47. (Three Times Amended) A method according to Claim 42 wherein the selectable marker is introduced into the pluripotent cell by targeted integration or random gene trap integration so as to be operatively coupled to a gene that is differentially expressed in neural progenitor cells.

48. (Three Times Amended) A method according to Claim 42 wherein the selectable marker is introduced into the pluripotent cell via random integration of a transgene in which the selectable marker is operatively coupled to a gene that is differentially expressed in neural progenitor cells.

49. (Three Times Amended) A method according to Claim 42 wherein the pluripotent cell is an ES, EG or EC cell and the method comprises forming an embryoid body in step (ii), or otherwise inducing differentiation of the cells.

54. (Amended) A method according to Claim 42 wherein the Sox gene is selected from Sox 1, Sox 2 and Sox 3.

65. (Amended) A method of generating purified neuron cells, comprising obtaining a culture of neural progenitor cells, using the method of Claim 42 wherein the selectable marker is differentially expressed in cells that express a Sox gene, and culturing the progenitor cells obtained in the presence of medium suitable for differentiation of the progenitor cells into neurons.

66. (New) A method for generating a culture that is purified or enriched in neural progenitor cells, comprising:

(i) introducing into a pluripotent cell a selectable marker that is differentially expressed in neural progenitor cells compared with its expression in other cells, wherein neural progenitor cells constitute a sub-set of the cells obtainable from the pluripotent cell, and wherein expression of the selectable marker is operatively linked to expression

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of a gene that is differentially expressed in neural progenitor cells,

(ii) culturing the pluripotent cell *in vitro* to induce differentiation of the pluripotent cell into a neural progenitor cell or into a mixture of cells including neural progenitor cells, or to induce preferential survival, in a mixed culture of cells, of neural progenitor cells; and

(iii) selecting for neural progenitor cells according to differential expression of the selectable marker introduced in step (i).

67. (New) A method according to Claim 66 wherein the pluripotent cell is selected from embryonic stem (ES) cells, embryonic germ (EG) cells, embryonic carcinoma (EC) cells, a primary culture of fetal cells, a primary culture of post-natal cells, and a primary culture of adult cells.

68. (New) A method according to Claim 66 comprising genetically modifying pluripotent cells by deleting, mutating, substituting, or adding genes in said pluripotent cells in order to assay gene function in neural progenitor cells, or render selected cells more suitable for transplantation, or both.

69. (New) A method according to Claim 66 further comprising:

(1) introducing into the pluripotent cell a second selectable marker that is differentially expressed in cells of a selectable sub-lineage compared with its expression in other cells, wherein cells of the selected sub-lineage are formed by differentiation of neural progenitor cells; and

(2) when a culture of neural progenitor cells has been obtained, allowing or inducing differentiation of the cells and selecting for cells that express the second selectable marker.

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70. (New) A method according to Claim 66 wherein the selectable marker is introduced into the pluripotent cell by targeted integration or random gene trap integration so as to be operatively coupled to a gene that is differentially expressed in neural progenitor cells.

71. (New) A method according to Claim 66 wherein the selectable marker is introduced into the pluripotent cell via random integration of a transgene in which the selectable marker is operatively coupled to a gene that is differentially expressed in neural progenitor cells.

72. (New) A method according to Claim 66 wherein the pluripotent cell is an ES, EG, or EC cell and the method comprises forming an embryoid body in step (ii), or otherwise inducing differentiation of the cells.

73. (New) A method according to Claim 72 wherein the differentiated cells are dissociated so as to form a culture substantially of individual cells.

74. (New) A method according to Claim 72 wherein differentiated cells of an embryoid body are dissociated using a protease, such as trypsin.

75. (New) A method according to Claim 66 wherein the selectable marker is an antibiotic resistance gene and the method comprises culture in the presence of antibiotic.

76. (New) A method according to Claim 66 wherein expression of the selectable marker is operatively linked to expression of a gene selected from the group consisting of Pax 3, delta-1, Mash-1, Math-4a, Pax 6,  $\beta$ 3-tubulin, synapsin-1, MAP2/tau, GFAP, GABA, and islet -1/2.

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